1/22

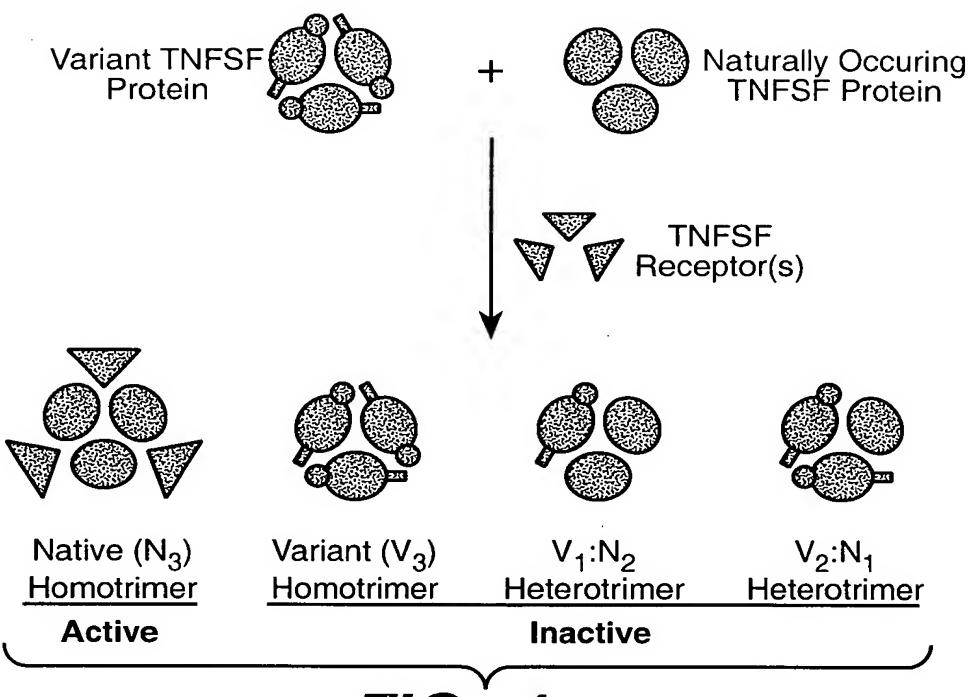


FIG._1



FIG._2

Figure 3

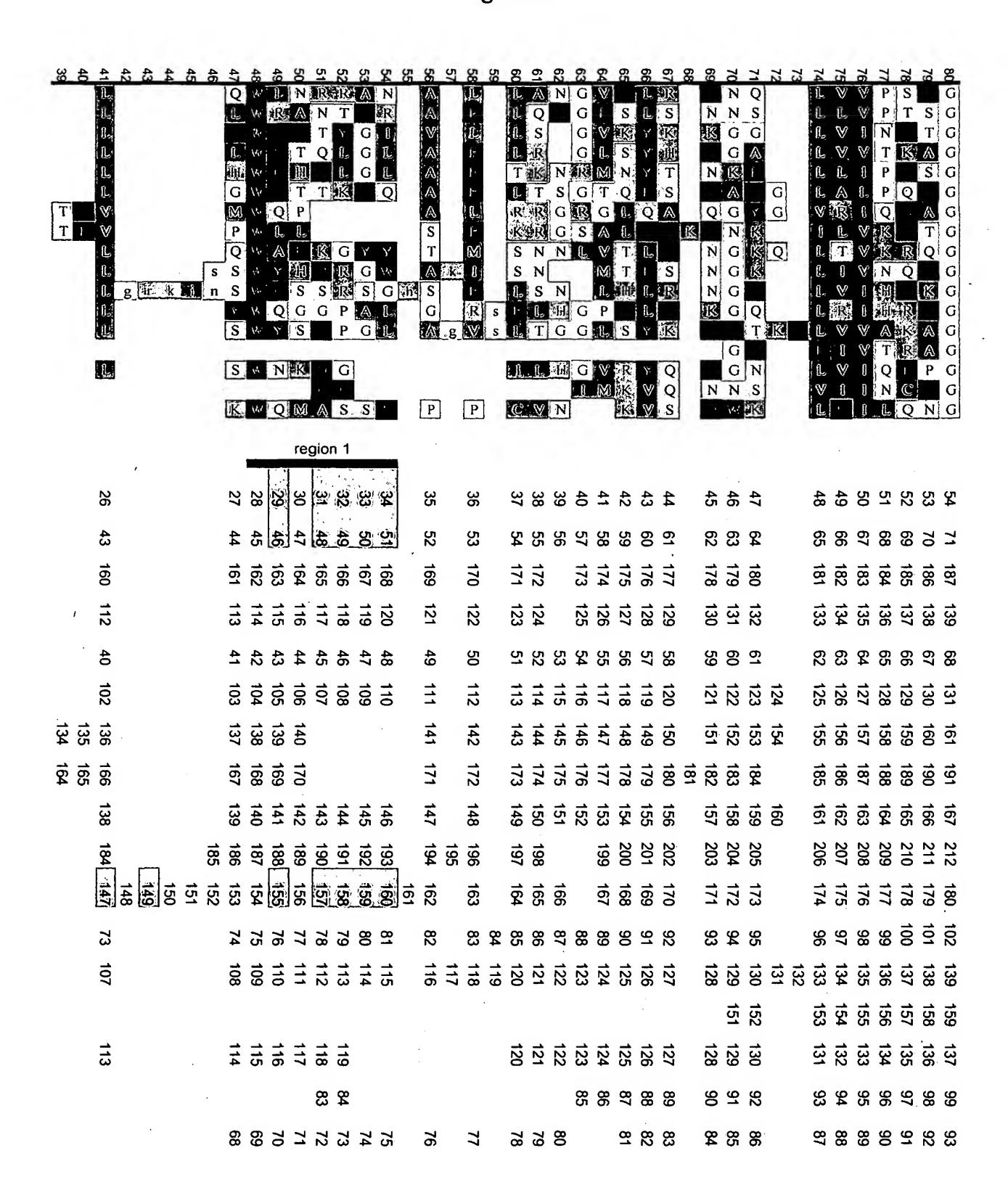


Figure 3

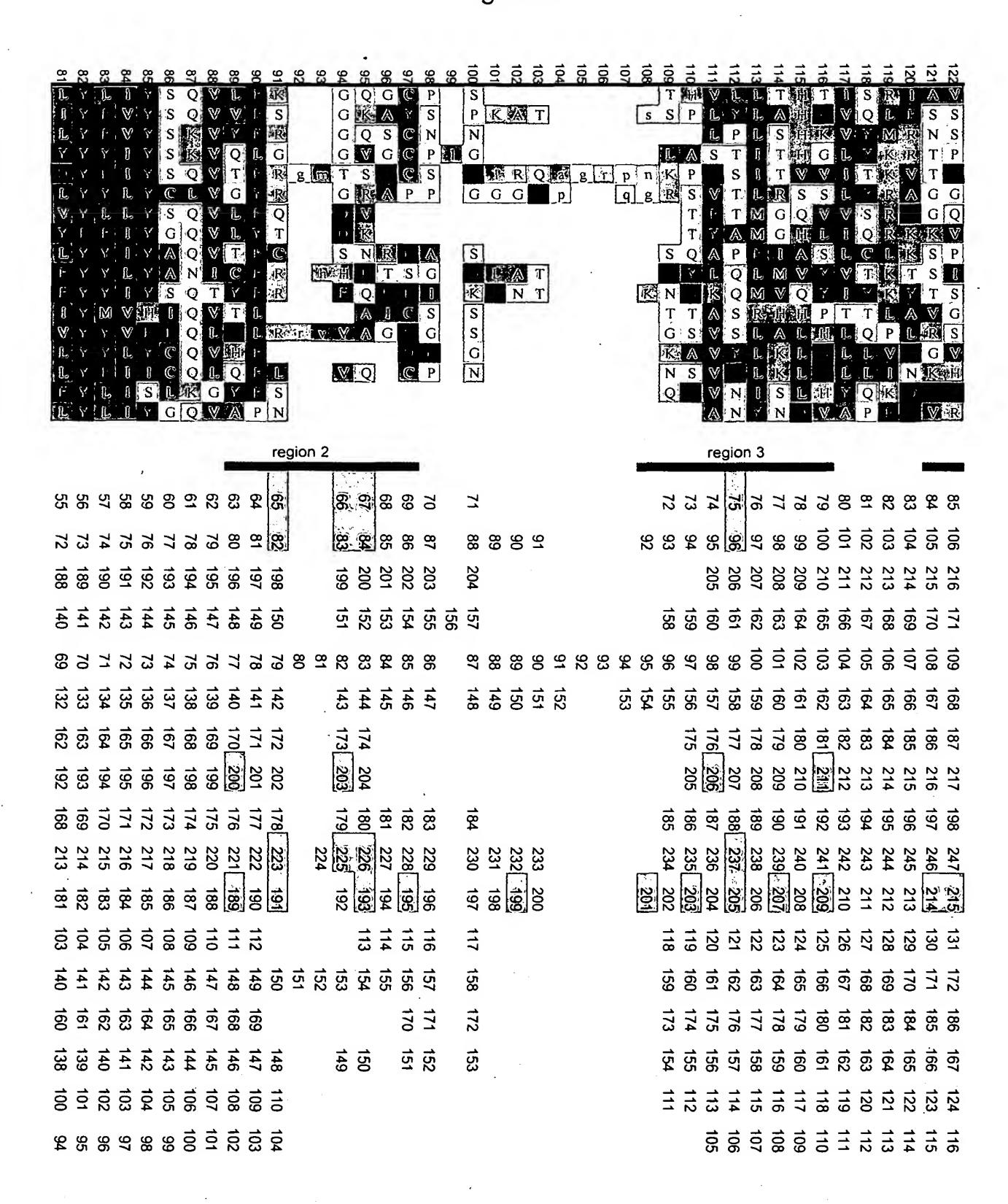


Figure 3

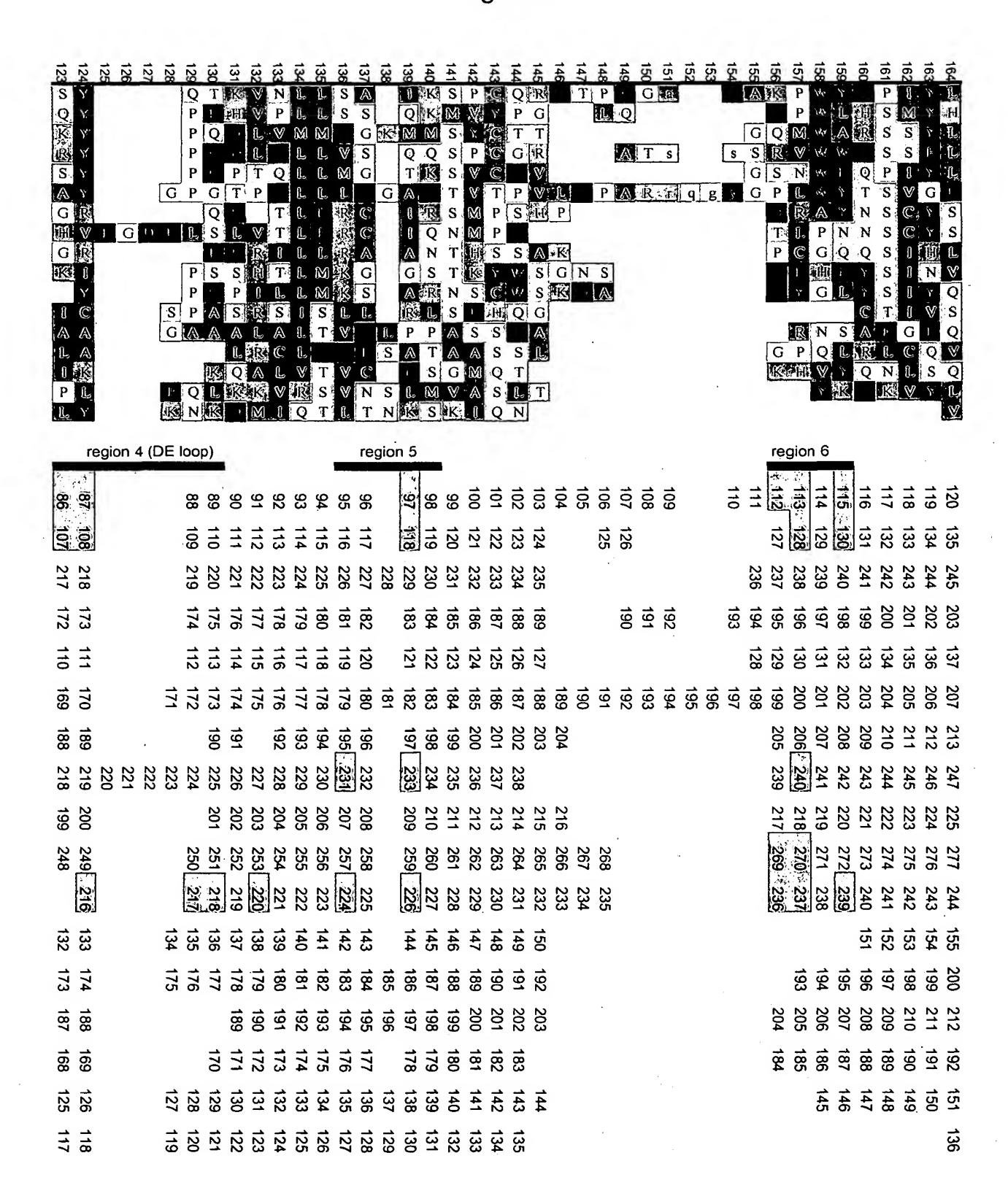


Figure 3

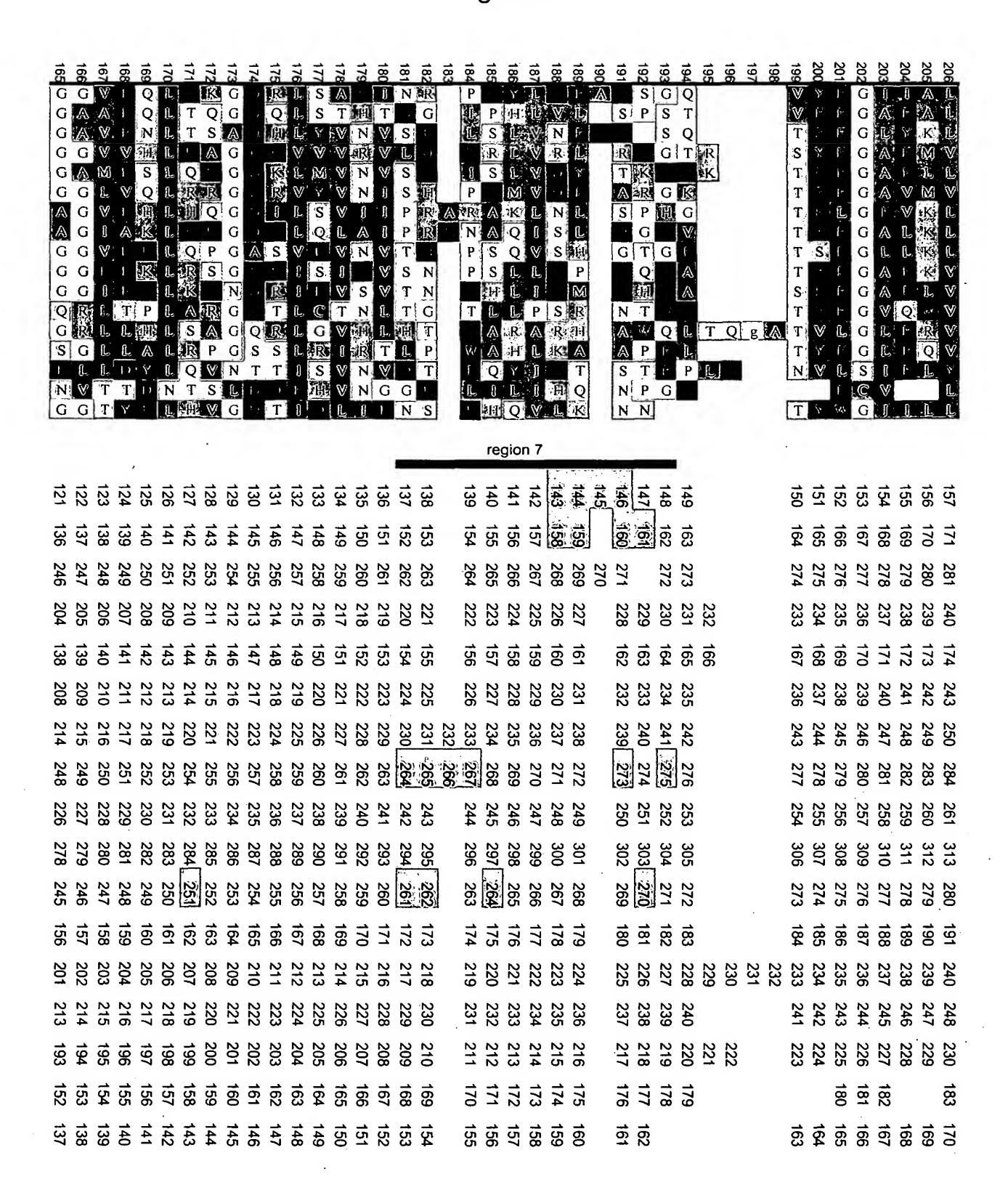
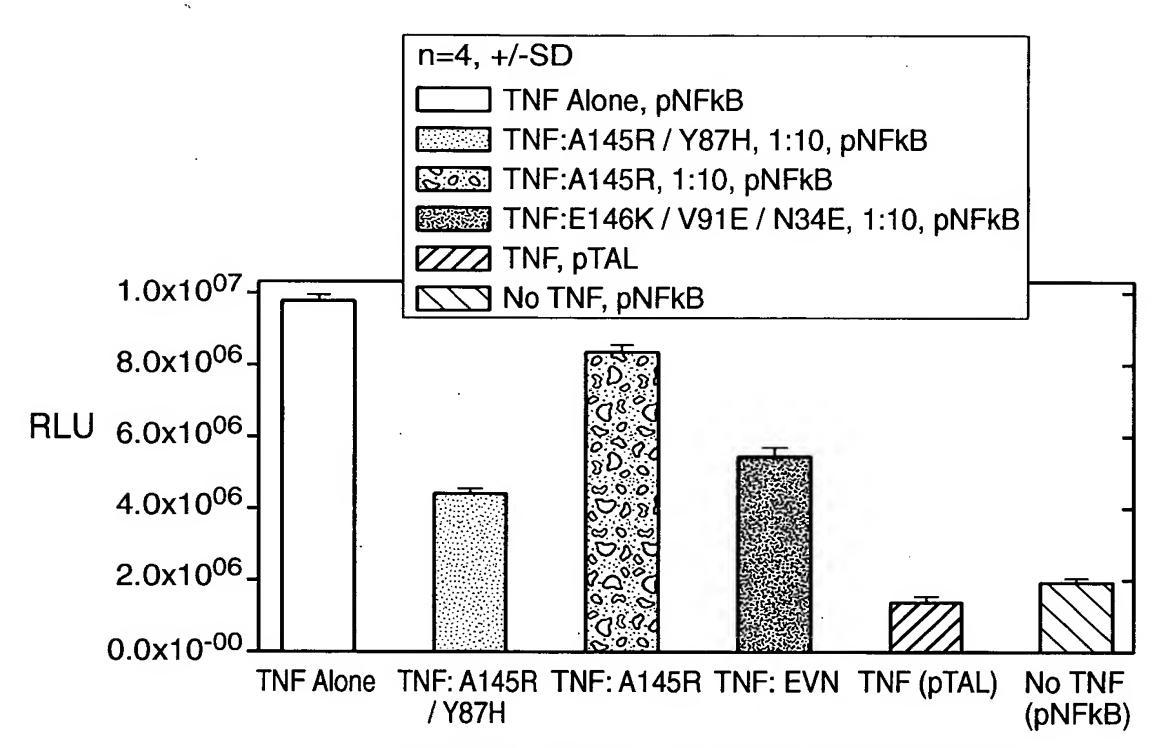


Figure 3

TNFSF Ligan		7 8 9 10 11 12 13 14 15 16 17 18 19 22 22 2 10 11 12 13 14 15 16 17 18 19 22 22	
TNFA TNFB (LT-α) FASL LIGHT	P R R	VARVVAN AARLTGK AARLTGA	P Q A G Q P S K Q N S S N S R S M P N S S L T g S G G P
VEGI TNFC (LT-β) APRIL BLyS	N Q L P S Q		P L KG QG NATSK S S V S PT D QKGS
CD40L RANKL TRAIL	Q D Q P Q R	AAHVIS I'AHLLTIN at VAAHITGTrgrsntis	A SS KTT SY PSGSM KVS sPNSKN KA
CD27L 4-1B8L	G P R Q G M	QQLVAQ	P R G P
TWEAK CD30L OX40L	A P KKKS	A P L Q V A K	IN KTR
AITRL		C M A K	G P L P S
,			
TNFA	1 12	13 15 15 15 15 15 15 15 15 15 15 15 15 15	25 24 22 20
TNFB (LT-α)	29	3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3	39 39 37
FASL	145	152 151 150 149 148 147	159 158 157 157 153
LIGHT	94	101 100 98 98 95	103
VEGI	36 35	37	36 36
TNFC (LT-β)	88 87	89 92 93 89 92 93 94 95	101 100 98 98
APRIL	116	123 123 122 121 120 119 118	133 132 133 131 130 128 127 128 127 126 127
BLyS	145 144	152 151 150 149 148 147	153 153 153 153
CD40L	122	129 128 127 126 125 124	137 138 139 130
RANKL	163	174 173 173 177 170 169 168 166 165	183 183 182 181 178 178 176
TRAIL	121	137 138 138 139 139 139 139 139 139 139 139 139 139	138 140 138
CD27L	67	68 69	71 72
4-1BBL	91 90 88	96 97 98 98	99 102 103 106
TWEAK	0		
CD30L	98 97 98 94 93	20 1 20 1 20 1 20 2 2 2 2 2 2 2 2 2 2 2	112 111 110 109 108
OX40L	7 4 73	81 80 79 76	8 2
AITRL		59 58	62 63 65 65 67



[10 ng/ml TNF α + 10-Fold Excess Variant]

FIG._4A

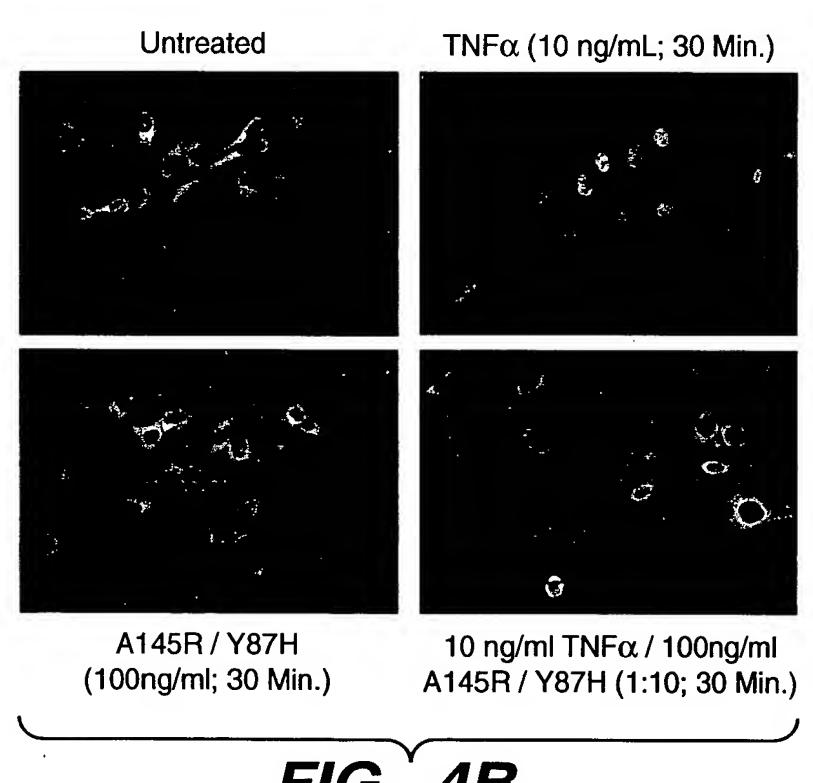
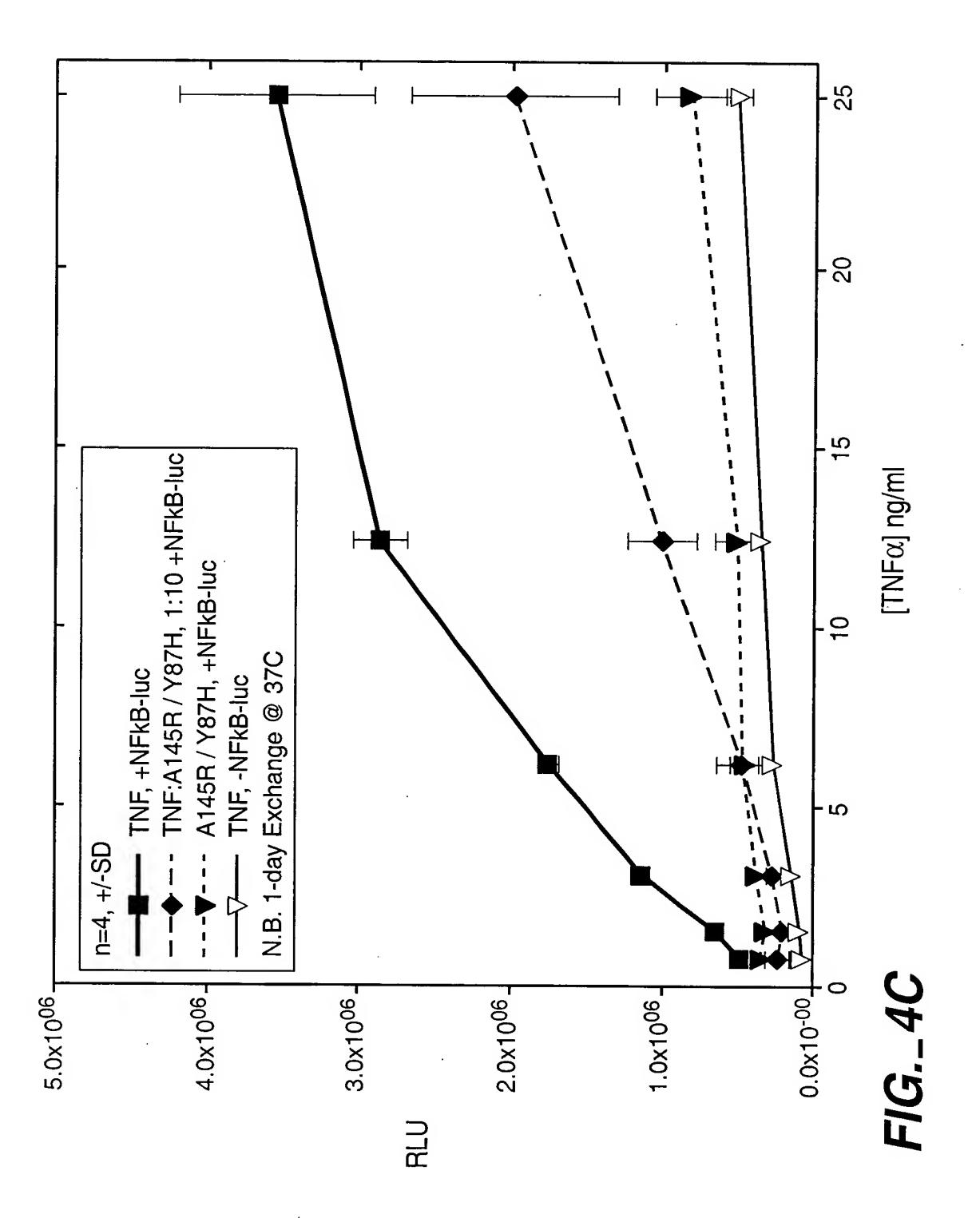
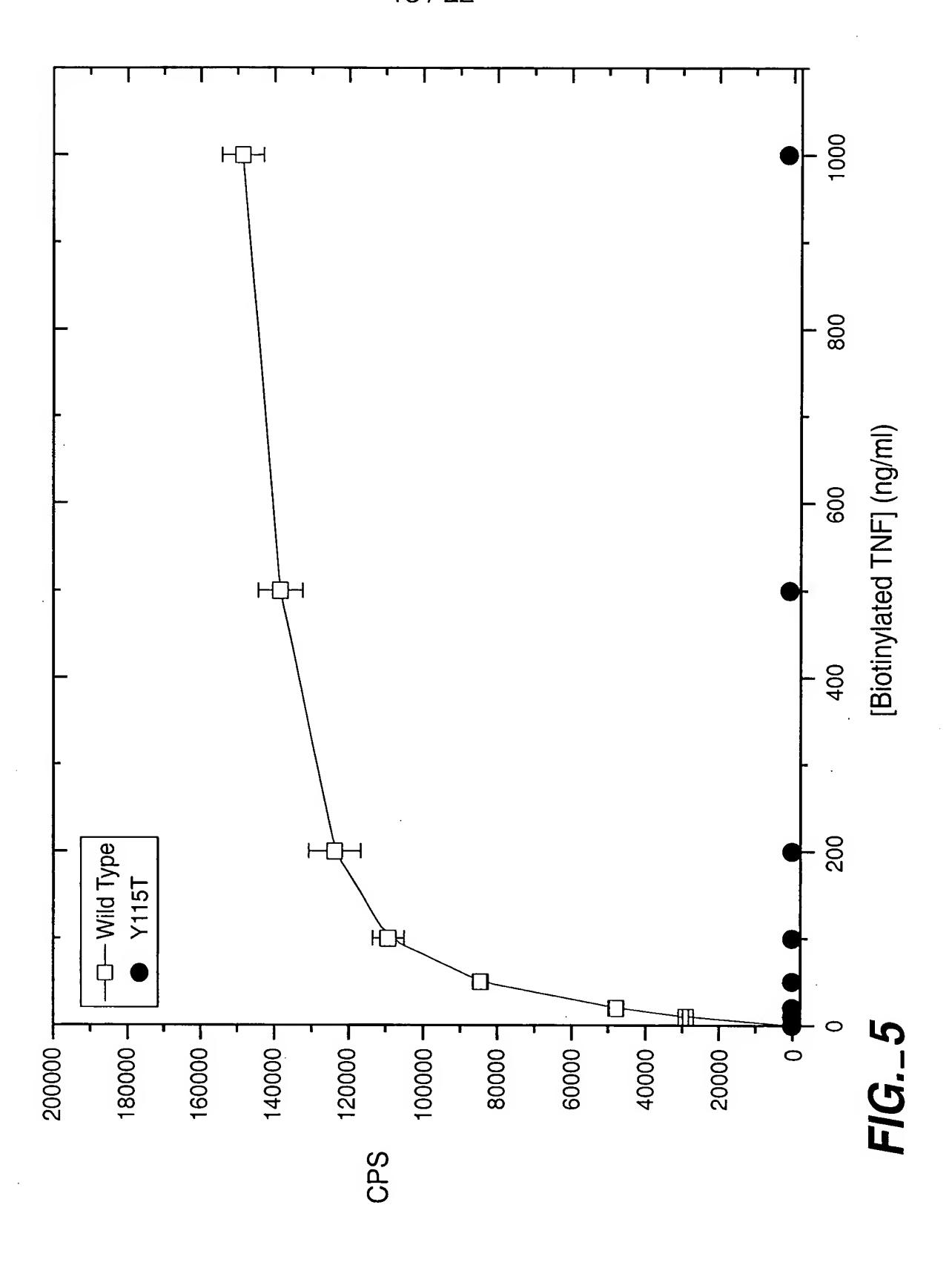


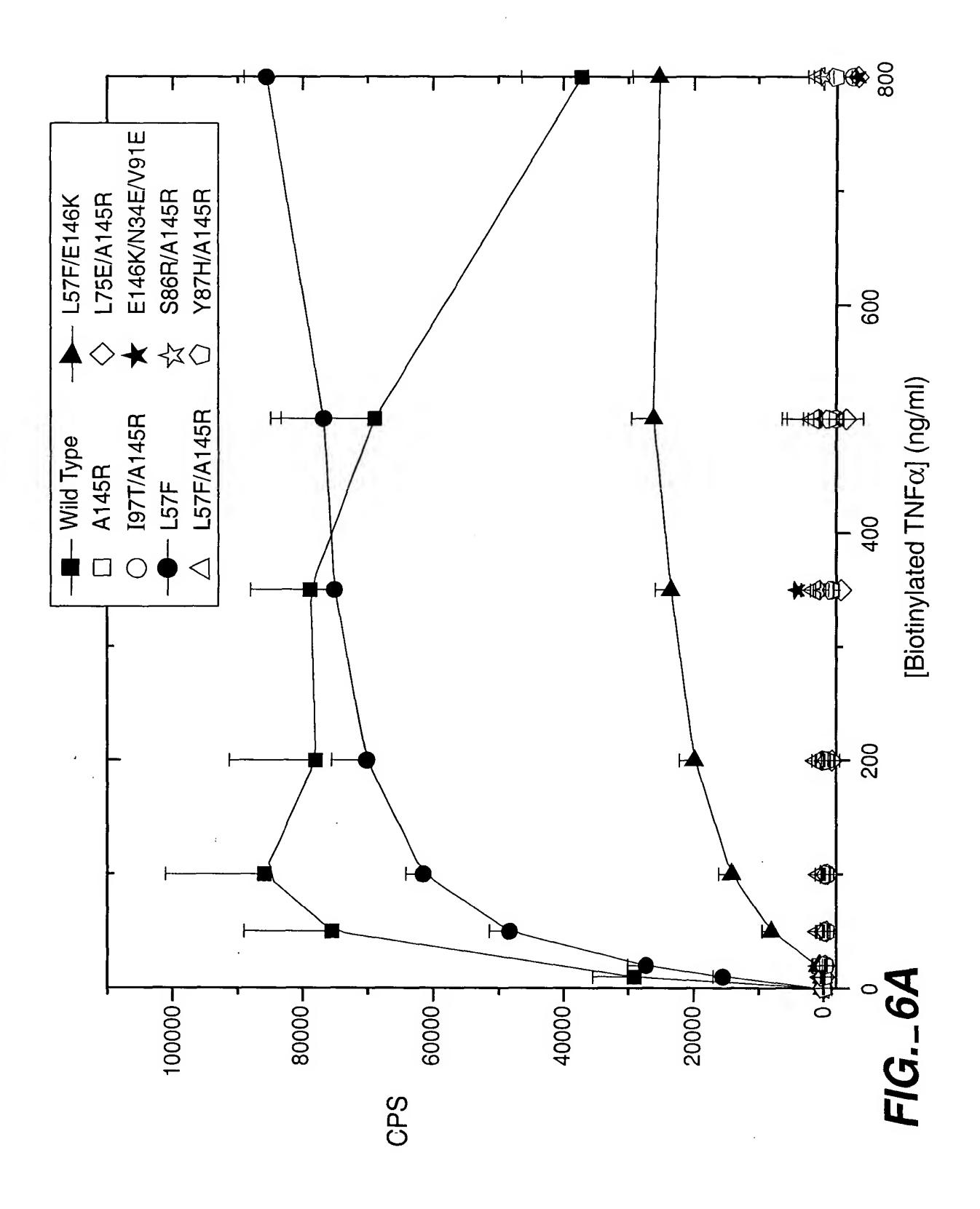
FIG. 4B



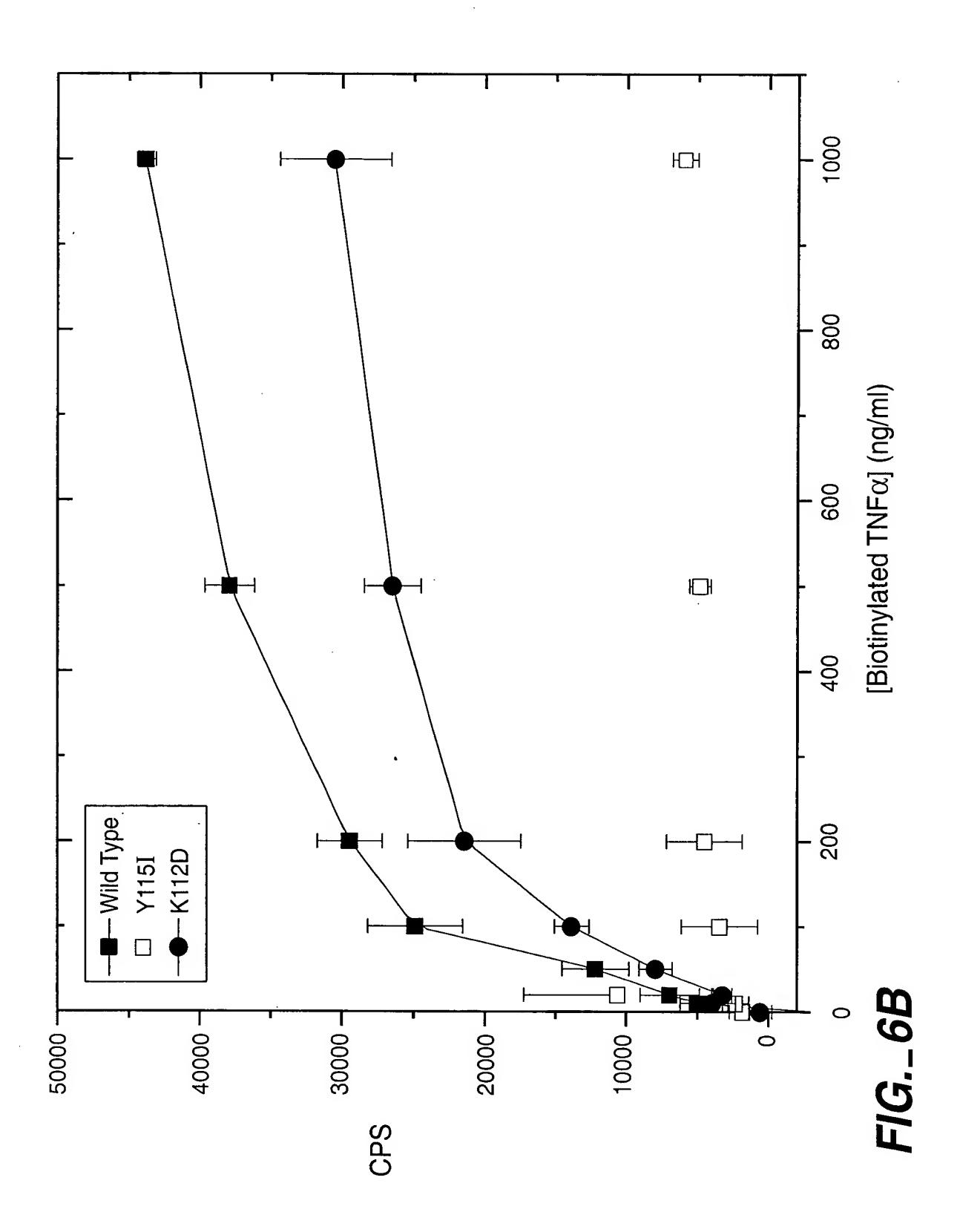
-

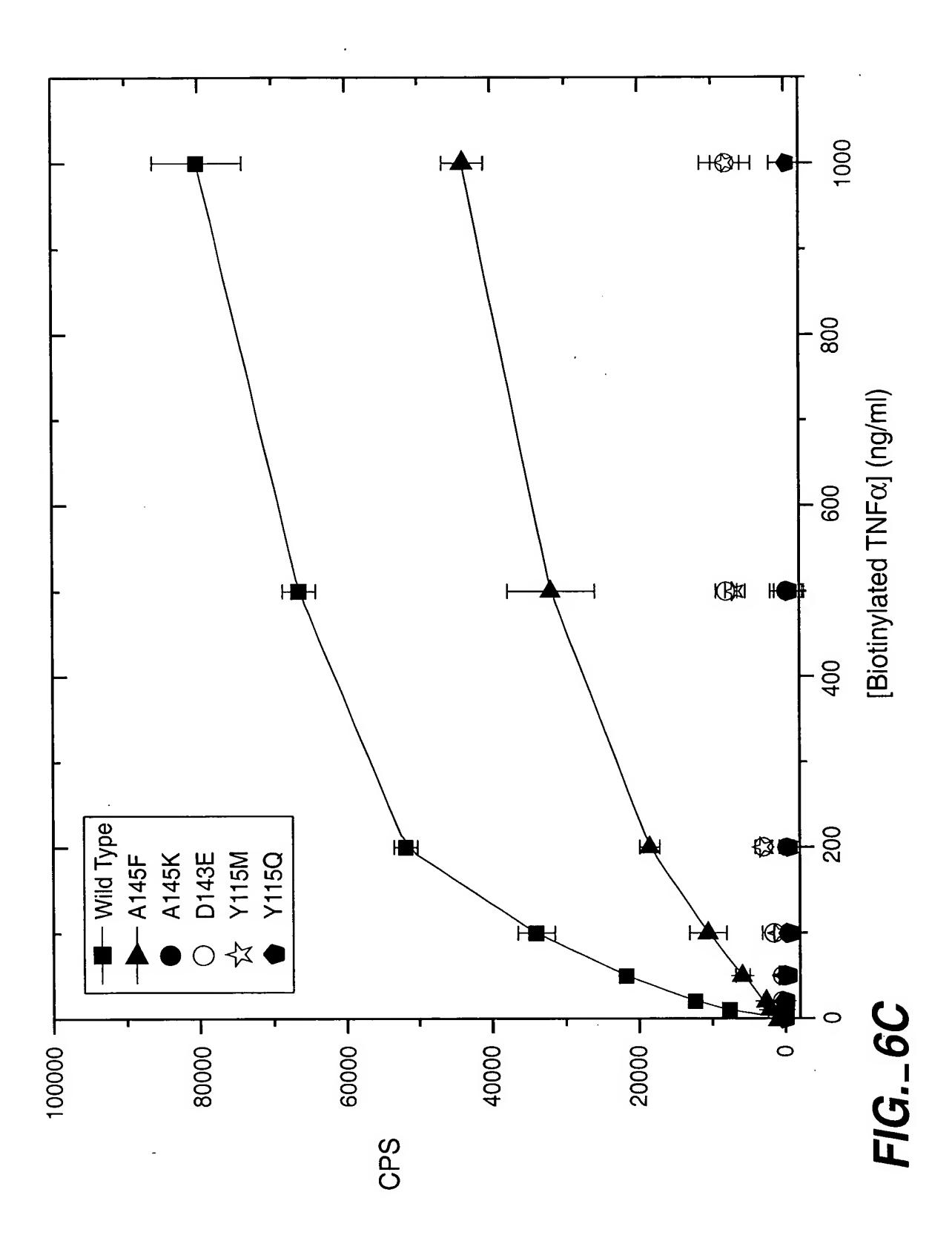


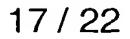
.



-







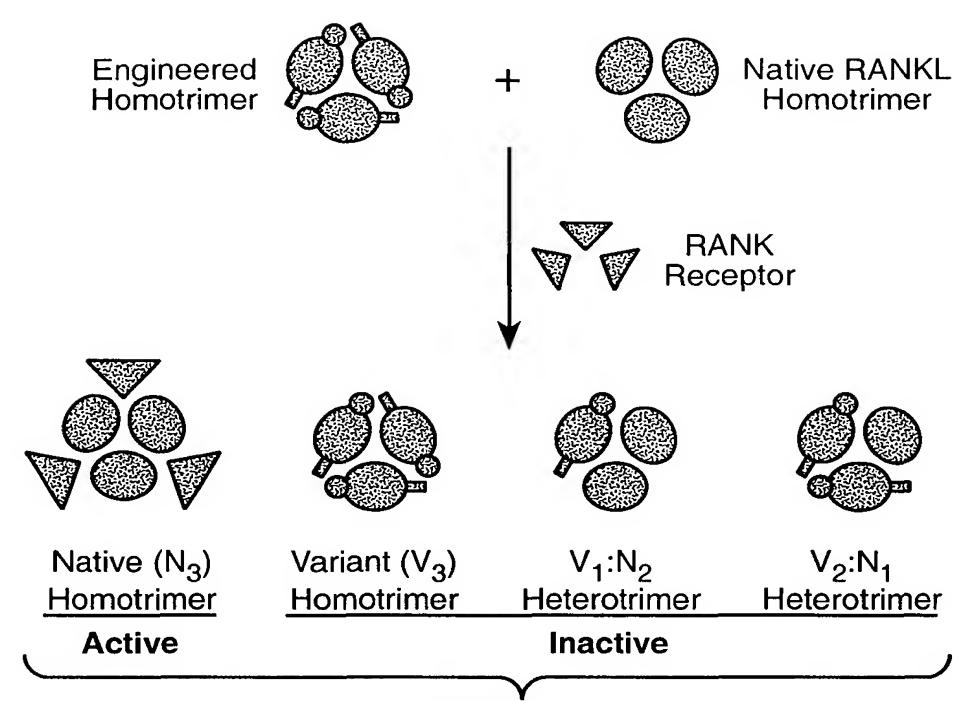
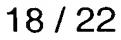
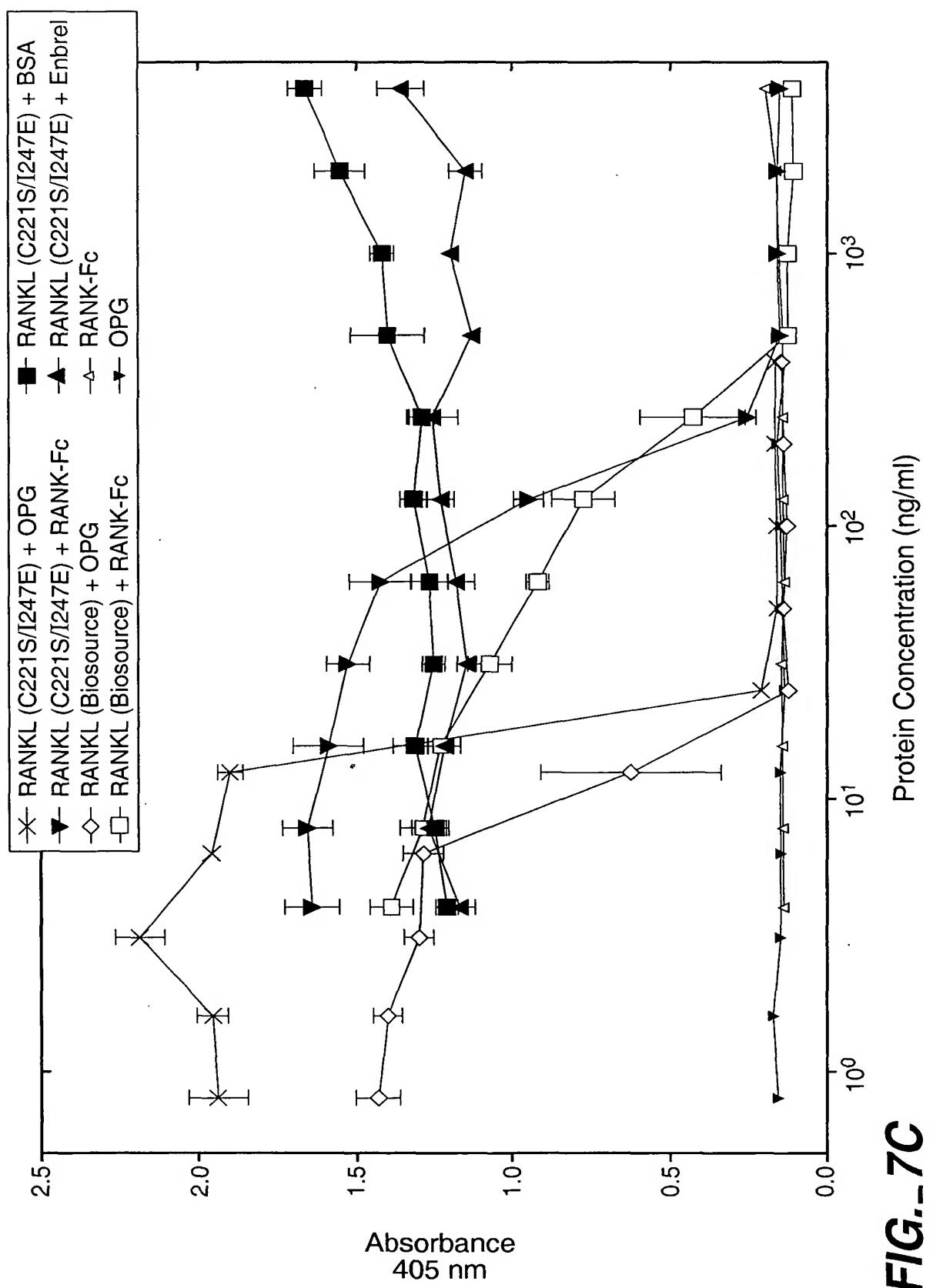


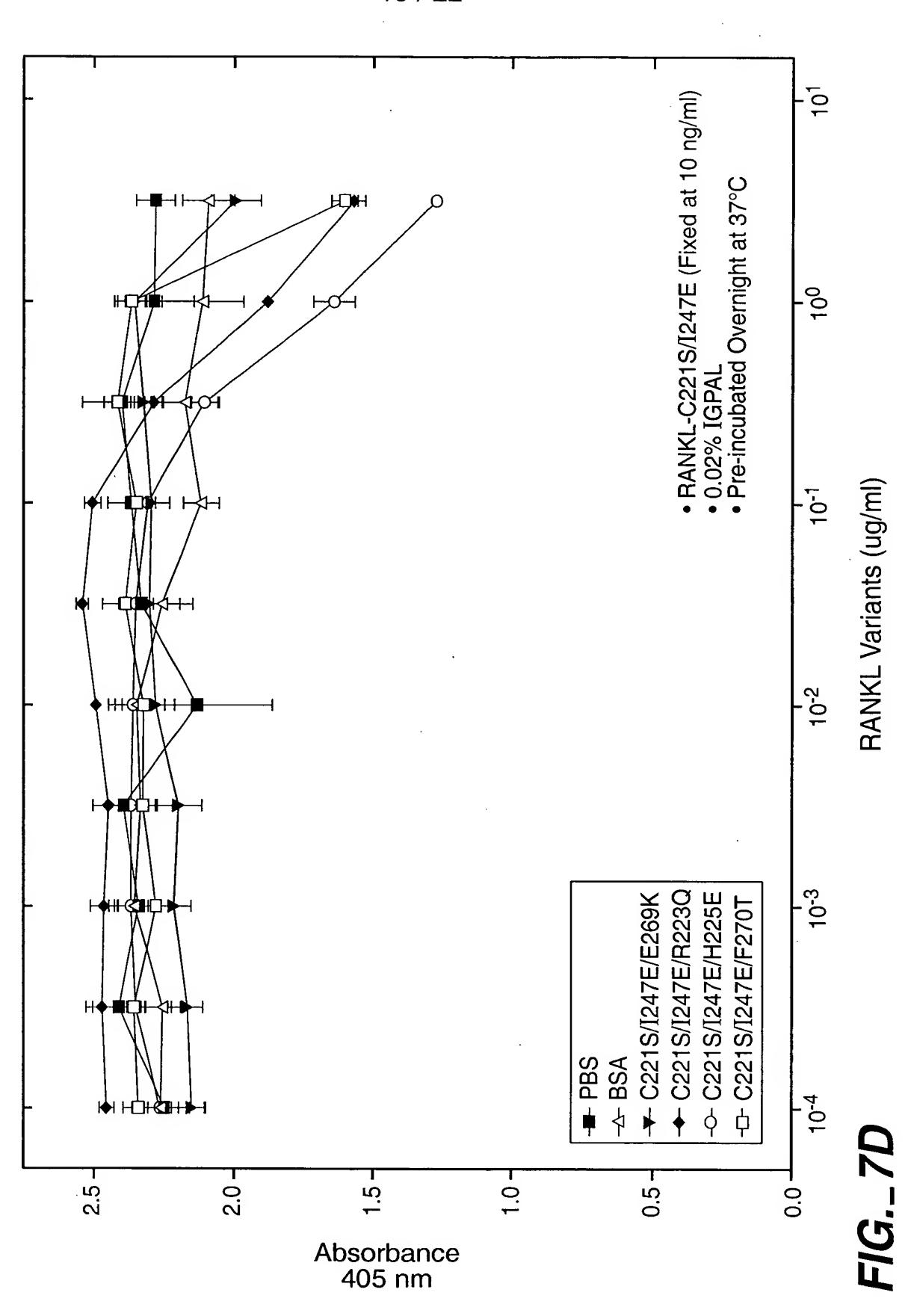
FIG. 7A

A172R	R191Q	K248E	1249R-S228E
A172Q	G191A	1249Q	K248E-T227K
S179E	R223Q	1249R	K248E-T227Q
S179D	R223E	H253T	R191K-I249R
H180E	R223M	H253S	R191E-I249R
H180N	H225R	S268D	G192A-K248E
H180S	H225N	E269R	D190Q-K248E
K181Q	H225T	E269Q	D190T-K248E
K181E	H225E	E269T	H225R-1249R
K181R	E226Q	E269K	H225N-1249R
S183K	E226R	F270V	E226Q-I249R
S183E	E226D	F270E	E226R-1249R
Y188F	T227K	F270T	H225R-G192A
Y188M	T227Q	F270K	H225N-G192A
Y188Q	S228H	R284E	H225R-D190Q
Y188E	S228E	S297Q	H225R-D190T
H189E	S228D	S297E	G192A-H225R-I249R
H189A	T233R	S297D	G192A-H225N-I249R
H189R	T233D	D300N	R191K-E226Q-K248E
D190Q	T233E	D302E	R191K-E226R-K248E
D190T	Q237K	D302Q	
R191K	Q237E	D302H	
R191E	Q237T	1249R-S228H	

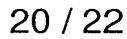
FIG._7B

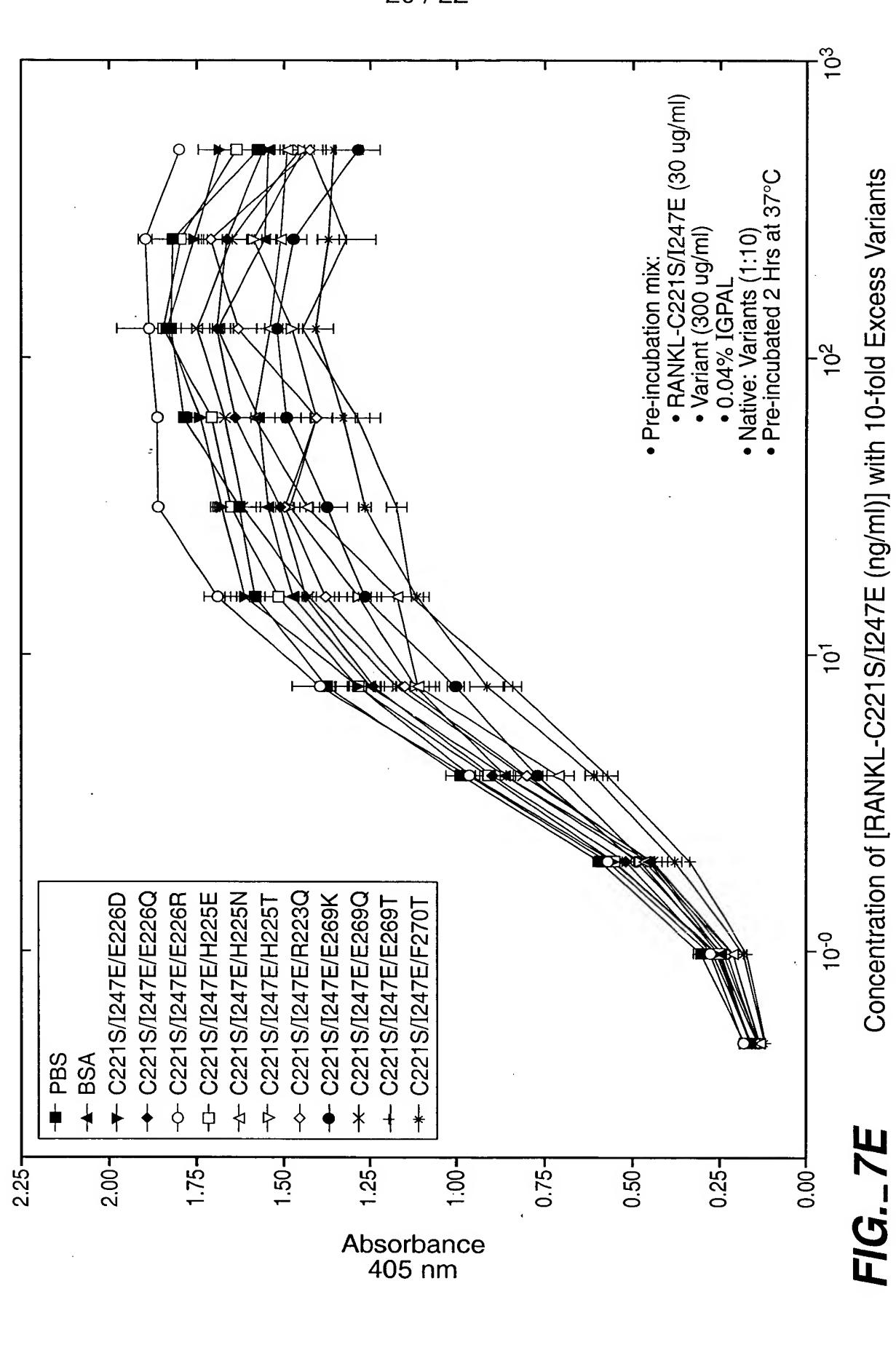


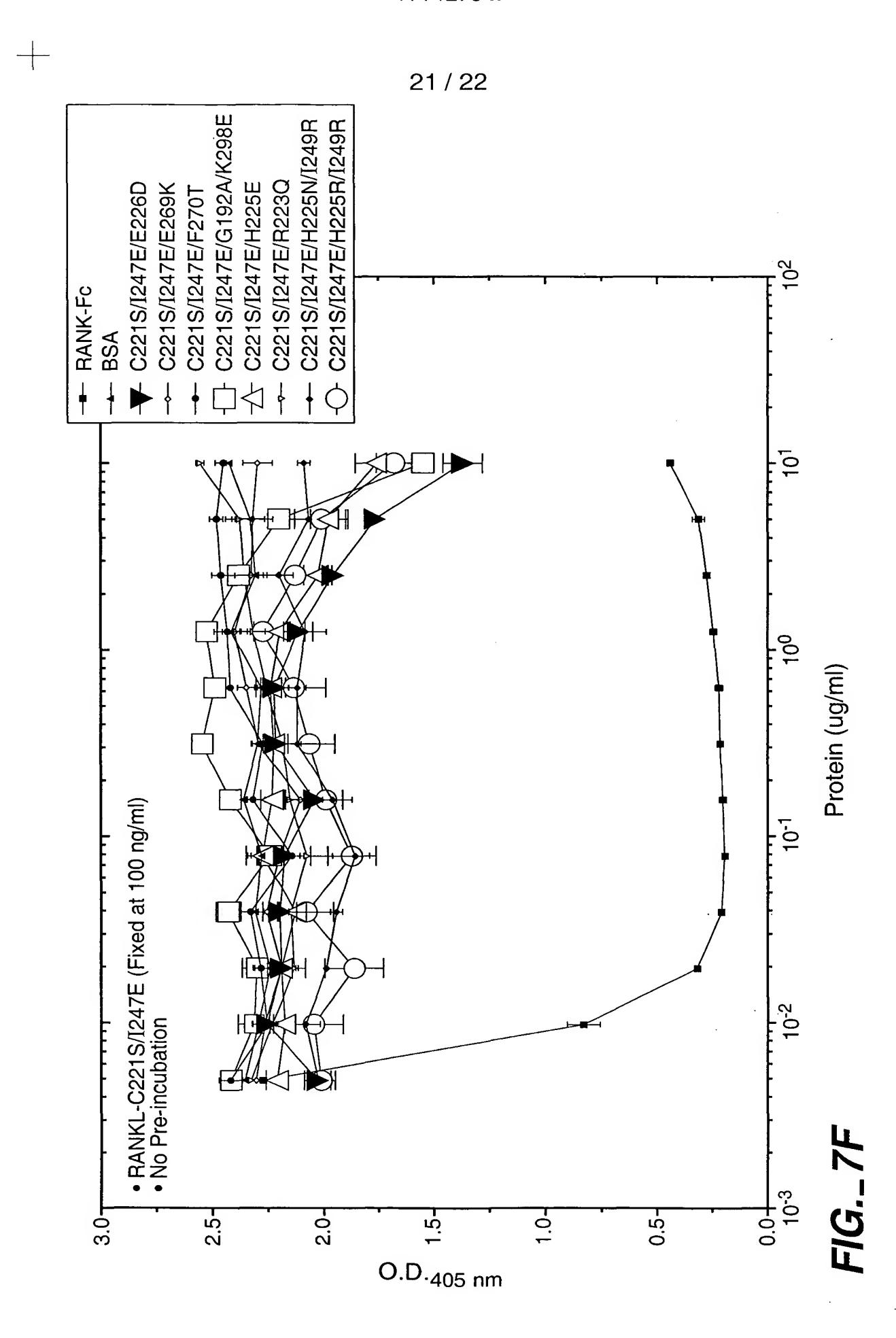




_-







C221S/I247E/E269T C221S/I247E/F270T C221S/I247E/E269K C221S/I247E/H225N C221S/I247E/H225E C221S/I247E/R223Q C221S/I247E/E226D C221S/I247E/G192A/K298E C221S/I247E/H225R/I249R

FIG._7G